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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Publication of

Christian GUELLY et al.

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Serial No. 10/528,031

Attorney Docket No. 2005_0070A

Filed March 16, 2005

POLYPEPTIDES AND NUCLEIC ACIDS
ENCODING THESE AND THEIR USE FOR
THE PREVENTION, DIAGNOSIS OR
TREATMENT OF LIVER DISORDERS
AND EPITHELIAL CANCER

[Corresponding to PCT/EP2003/010564
Filed September 23, 2003]

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT

Commissioner for Patents
P.O. Box 1450
Alexandria, VA 22313-1450

Sir:

Pursuant to the provisions of 37 CFR 1.56, 1.97 and 1.98, Applicants request consideration of the references listed on attached form PTO-1449 and any additional information identified below in paragraph 3. A legible copy of each reference listed on the Form PTO-1449 is enclosed, except a copy is not provided for:

- ☒ each U.S. Patent and U.S. Patent application publication;
- ☐ each reference previously cited in the international application
PCT/_____; and/or
- ☐ each reference previously cited in prior parent application Serial No.
_____.

1a. ☒ This Information Disclosure Statement is submitted:

within three months of the filing date (or of entry into the National Stage) of the above-entitled application, **or**

before the mailing of a first Office Action on the merits or the mailing of a first Office Action after the filing of an RCE,

and thus no certification and/or fee is required.

1b. ☐ This Information Disclosure Statement is submitted

after the events of above paragraph 1a and prior to the mailing date of a final Office Action or a Notice of Allowance or an action which otherwise closes prosecution in the application, and thus:

(1) ☐ the certification of paragraph 2 below is provided, **or**

(2) ☐ the fee of \$180.00 specified in 37 CFR 1.17(p) is enclosed.

1c. ☐ This Information Disclosure Statement is submitted:

after the mailing date of a final Office Action or Notice of Allowance or action which otherwise closes prosecution in the application, and prior to payment of the issue fee, and thus:

the certification of paragraph 2 below is provided, and

the fee of \$180.00 specified in 37 CFR 1.17(p) is enclosed.

2. It is hereby certified

a. ☐ that each item of information contained in this Information Disclosure Statement was first cited in any communication from a foreign patent office in a counterpart foreign application not more than three months prior to the filing of the Statement, or

b. ☐ that no item of information contained in the Information Disclosure Statement was cited in a communication from a foreign patent office in a counterpart foreign application and, to the knowledge of the person signing the certification

after making reasonable inquiry, was known to any individual designated in §1.56(c) more than three months prior to the filing of the Statement.

3. ☐ Consideration of the following list of additional information (including any copending or abandoned U.S. application, prior uses and/or sales, etc.) is requested.
4. For each non-English language reference listed on the attached form PTO-1449, reference is made to:
- a. ☐ a full or partial English language translation submitted herewith,
 - b. ☐ a foreign patent office search report (in the English language) submitted herewith,
 - c. ☐ the concise explanation contained in the specification of the present application at page,
 - d. ☐ the concise explanation set forth in the attached English language abstract,
 - e. ☐ the concise explanation set forth below or on a separate sheet attached to the reference:
5. ☐ A foreign patent office search report citing one or more of the references is enclosed.

Respectfully submitted,

Christian GUELLY et al.

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July 7, 2005

INFORMATION DISCLOSURE STATEMENT

FORM PTO 1449 (modified)

U.S. DEPARTMENT OF COMMERCE
PATENT AND TRADEMARK OFFICELIST OF REFERENCES CITED BY APPLICANT(S)
(Use several sheets if necessary)

Date Submitted to PTO: July 7, 2005

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GROUP

U.S. PATENT DOCUMENTS

| *EXAMINER INITIAL | | DOCUMENT NUMBER | DATE | NAME | CLASS | SUBCLASS | FILING DATE IF APPROPRIATE |
|----------------------|----|--------------------|---------|----------------|-------|----------|-------------------------------|
| | AA | 4,736,866 | 4/1988 | Leder et al. | | | |
| | AB | 4,816,397 | 3/1989 | Boss et al. | | | |
| | AC | 4,816,567 | 3/1989 | Cabilly et al. | | | |
| | AD | 5,569,588 | 10/1996 | Ashby et al. | | | |
| | AE | 5,583,278 | 12/1996 | Alt et al. | | | |
| | AF | 5,625,122 | 4/1997 | Mak | | | |
| | AG | 5,698,765 | 12/1997 | Mak | | | |
| | AH | 5,744,305 | 4/1998 | Fodor et al. | | | |
| | AI | 5,750,825 | 5/1998 | Yazaki et al. | | | |

FOREIGN PATENT DOCUMENTS

| | | DOCUMENT NUMBER | DATE | COUNTRY | CLASS | SUBCLASS | TRANSLATION YES NO | |
|--|----|--------------------|---------|---------|-------|----------|-----------------------|--|
| | AJ | 196 19 373 | 11/1997 | DE | | | Abstract | |
| | AK | 196 25 049 | 1/1998 | DE | | | Abstract | |
| | AL | 198 16 414 | 10/1999 | DE | | | Abstract | |
| | AM | 0 127 839 | 12/1984 | EP | | | | |
| | AN | 0 368 684 B1 & B2 | 5/1990 | EP | | | | |

OTHER DOCUMENT(S) (Including Author, Title, Date, Pertinent Pages, Etc.)

| | | |
|--|----|---|
| | AO | B. Alberts et al., eds., The Molecular Biology of the Cell , 3 rd Edition, 2002 Garland, New York, USA, pp. 98-137. |
| | AP | R. N. M. MacSween et al., eds., Pathology of the Liver, 4 th Edition, 2001, Churchill Livingstone, Edinburgh, pp. 598, 732. |
| | AQ | M. Hollander et al., eds., Nonparametric Statistical Methods, 2 nd Edition, 1973 New York, John Wiley & Sons, pp. 26-33 and 68-75. |

EXAMINER

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|----------------------|----|--------------------|--------|-----------------|-------|----------|-------------------------------|
| | BA | 6,183,956 | 2/2001 | Sivaraja et al. | | | |
| | BB | | | | | | |

FOREIGN PATENT DOCUMENTS

| | | DOCUMENT NUMBER | DATE | COUNTRY | CLASS | SUBCLASS | TRANSLATION YES NO | |
|--|----|--------------------|--------|---------|-------|----------|-----------------------|--|
| | BC | 88/01649 | 3/1988 | WO | | | | |
| | BD | 93/06213 | 4/1993 | WO | | | | |
| | BE | 95/11910 | 5/1995 | WO | | | | |
| | BF | 97/29116 | 8/1997 | WO | | | | |
| | BG | 98/24884 | 6/1998 | WO | | | | |
| | BH | 98/37240 | 8/1998 | WO | | | | |

OTHER DOCUMENT(S) (Including Author, Title, Date, Pertinent Pages, Etc.)

| | | |
|--|----|--|
| | BI | M. Kozak, "An analysis of 5'-noncoding sequences 699 vertebrate messenger RNAs", Nucleic Acids Research, Vol. 15, No. 20, pp. 8125-8148, 1987. |
| | BJ | S. Kawata et al., "Effect of pravastatin on survival in patients with advanced hepatocellular carcinoma. A randomized controlled trial", British Journal of Cancer, Vol. 84, No. 7, pp. 886-891, 2001. |
| | BK | C. K. Shortell et al., "Hepatic Adenoma and Focal Nodular Hyperplasia", Surgery, Gynecology & Obstetrics, Vol. 173, pp. 426-431, 1991. |
| | BL | W. Lau et al., "Differential Gene Expression of Hepatocellular Carcinoma using cDNA Microarray Analysis", Oncology Research, Vol. 12, pp. 59-69, 2000. |
| | BM | M. A. Buendia et al., "Genetic Alterations in Hepatoblastoma and Hepatocellular Carcinoma: Common and Distinctive Aspects", Med. Pediatr. Oncol., Vol. 39, No. 5, pp. 530-535, November 2002. |
| | BN | M. Mencinger et al., "Expression analysis and chromosomal mapping of a novel human gene, APRIL, encoding an acidic protein rich in leucines", Biochimica et Biophysica Acta, Vol. 1395, pp. 176-180, 1998. |

EXAMINER

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| | CA | | | | | | |

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| | CB | | | | | | |

OTHER DOCUMENT(S) (Including Author, Title, Date, Pertinent Pages, Etc.)

| | | |
|--|----|---|
| | CC | D. F. Bauer, "Constructing Confidence Sets Using Rank Statistics", Journal of the American Statistical Association, Vol. 67, No. 339, pp. 687-690, September 1972. |
| | CD | R. Q. H. Zheng et al., "Inhibition of gene expression by anti-sense oligodeoxynucleotides", Clin. Exp. Immunol., Vol. 100, pp. 380-382, 1995. |
| | CE | W. Nellen et al., "What makes an mRNA anti-sense-itive", Trends Biochem., Sci., Vol. 18, pp. 419-423, November 18, 1993. |
| | CF | M. Amarzguoui et al., "Hammerhead ribozyme design and application", Cellular and Molecular Life Sciences, Vol. 54, pp. 1175-1202, 1998. |
| | CG | L. Couture et al., "Anti-gene therapy: the use of ribozymes to inhibit gene function", Trends Genet., Vol. 12, No. 12, pp. 510-515, December 1996. |
| | CH | E. Uhlmann et al., "Antisense Oligonucleotides: A New Therapeutic Principle", Chemical Reviews, Vol. 90, No. 4, pp. 543-584, June 1990. |
| | CI | M. Gossen et al., "Inducible gene expression systems for higher eukaryotic cells", Curr., Opin. Biotechnol., Vol. 5, pp. 516-520, 1994. |
| | CJ | H. Schrem et al., "Liver-Enriched Transcription Factors in Liver Function and Development. Part I: The Hepatocyte Nuclear Factor Network and Liver-Specific Gene Expression", Pharmacological Reviews, Vol. 54, No. 1, pp. 129-158, 2002. |
| | CK | M. Pontoglio et al., "Hepatocyte Nuclear Factor 1 α Controls the Expression of Terminal Complement Genes", J. Exp. Med., Vol. 194, No. 11, pp. 1683-1689, December 3, 2001. |
| | CL | J. Zwicker et al., "Cell-cycle regulation of gene expression by transcriptional repression", Trends Genet., Vol. 13, No. 1, pp. 3-6, January 1997. |

EXAMINER

DATE CONSIDERED

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|--|----|--------------------|------|---------|-------|----------|-----------------------|
| | DB | | | | | | |
| | DC | | | | | | |

OTHER DOCUMENT(S) (Including Author, Title, Date, Pertinent Pages, Etc.)

| | | |
|--|----|--|
| | DD | D. Lindemann et al., "Versatile Retrovirus Vector Systems for Regulated Gene Expression <i>In Vitro</i> and <i>In Vivo</i> ", Molecular Medicine, Vol. 3, No. 7, pp. 466-476, July 1997. |
| | DE | M. L. Springer et al., "VEGF Gene Delivery to Muscle: Potential Role for Vasculogenesis in Adults", Molecular Cell, Vol. 2, pp. 549-558, November 1998. |
| | DF | Polites and Pinkert, Transgenic Animal Technology, Chapter 2: DNA Microinjection and Transgenic Animal Production, pp. 15-68, 1994. |
| | DG | T. Doetschman, Transgenic Animal Technology, Chapter 4: Gene Targeting in Embryonic Stem Cells: I. History and Methodology", pp. 113-141, 1994. |
| | DH | E. Harlow et al., eds., Antibodies, A Laboratory Manual, Chapter 5: Immunizations, Cold Spring Harbor Press, New York, USA, pp. 53-135, 1988. |
| | DI | A. Knappik et al., "Fully Synthetic Human Combinatorial Antibody Libraries (HuCAL) Based on Modular Consensus Frameworks and CDRs Randomized with Trinucleotides", J. Mol. Biol., Vol. 296, pp. 57-86, 2000. |
| | DJ | K. G. Ishak et al., "Tumors of the Liver and Intrahepatic Bile Ducts", Atlas of Tumor Pathology, Armed Forces Institute of Pathology, Wash. DC, pp. 199-231, 1999. |
| | DK | H. E. Chadd et al., "Therapeutic Antibody Expression Technology", Curr. Opin. Biotechnol., Vol. 12, pp. 188-194, 2001. |
| | DL | J. Sambrook et al., "Molecular Cloning", A Laboratory Manual, Second Edition, Construction and Analysis of cDNA Libraries, Chapter 8, pp. 8.1-8.81, 1989. |

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DATE CONSIDERED

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| | EB | | | | | | |

OTHER DOCUMENT(S) (Including Author, Title, Date, Pertinent Pages, Etc.)

| | | |
|--|----|--|
| | EC | J. Sambrook et al., "Molecular Cloning", A Laboratory Manual, Second Edition, Analysis and Cloning of Eukaryotic Genomic DNA, Chapter 9, pp. 9.47-9.58, 1989. |
| | ED | J. Sambrook et al., "Molecular Cloning", A Laboratory Manual, Second Edition, Preparation of Radiolabeled DNA and RNA Probes, Chapter 10, pp. 10.1-10.67, 1989. |
| | EE | S. Fields et al., "The two-hybrid system: an assay for protein-protein interactions", Trends in Genetics, Vol. 10, No. 8, pp.286-292, August 1994. |
| | EF | J. Sambrook et al., "Molecular Cloning", A Laboratory Manual, Second Edition, Enzymes used in Molecular Cloning, pp. 5.52-5.55, 1989. |
| | EG | J. Sambrook et al., "Molecular Cloning", A Laboratory Manual, Second Edition, Extraction, Purification, and Analysis of Messenger RNA from Eukaryotic Cells, pp. 7.39-7.52, 1989. |
| | EH | H. M. Cooper et al., "Preparation of Polyclonal Antisera", Current Protocols in Molecular Biology, Suppl. 50, pp. 11.12.1-11.12.9, 2000. |
| | EI | S. A. Fuller et al., "Preparation of Monoclonal Antibodies", Current Protocols in Molecular Biology, Supplement 18, pp. 11.4.1-11.9.3, 1992. |
| | EJ | R. K. Scopes et al., "Analysis of Proteins", Current Protocols in Molecular Biology, pp. 10.0.1-10.0.20, 1998. |
| | EK | D. Tackels-Horne et al., "Identification of Differentially Expressed Genes in Hepatocellular Carcinoma and Metastatic Liver Tumors by Oligonucleotide Expression Profiling", Cancer, Vol. 92, No. 2, pp. 395-405, July 15, 2001. |

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| | FB | | | | | | |

OTHER DOCUMENT(S) (Including Author, Title, Date, Pertinent Pages, Etc.)

| | | |
|--|----|--|
| | FC | A. Persidis, "Ribozyme Therapeutics", Nature Biotechnology, Vol. 15, pp. 921-922, September 1997. |
| | FD | M. Schena et al., "Quantitative Monitoring of Gene Expression Patterns with a Complementary DNA Microarray", Science, Vol. 270, pp. 467-470, October 1995. |
| | FE | L. Beigelman et al., "Synthesis and Biological Activities of a Phosphorodithioate Analog of 2',5'-oligoadenylate", Nucleic Acids Research, Vol. 23, No. 19, pp. 3989-3994, 1995. |
| | FF | D. J. Lockhart et al., "Expression monitoring by hybridization to high-density oligonucleotide arrays", Nature Biotechnology, Vol. 14, pp. 1675-1680, December 1996. |
| | FG | A. Blanchard, "Sequence to array: Probing the genome's secrets", Nature Biotechnology, Vol. 14, p. 1649, December 1996. |
| | FH | H. F. Kawai et al., "α-Fetoprotein-Producing Hepatoma Cell Lines Share Common Expression Profiles of Genes in Various Categories Demonstrated by cDNA Microarray Analysis", Hepatology, Vol. 33, No. 3, pp. 676-691, 2001. |
| | FI | H. Nagai et al., "Genomic Aberrations in Early Stage Human Hepatocellular Carcinomas", Cancer, Vol. 82, No. 3, pp. 454-461, February 1, 1998. |
| | FJ | H. Okabe et al., "Genome-wide Analysis of Gene Expression in Human Hepatocellular Carcinomas Using cDNA Microarray: Identification of Genes Involved in Viral Carcinogenesis and Tumor Progression", Cancer Research, Vol. 61, pp. 2129-2137, March 1, 2001. |
| | FK | M. Salvucci et al., "Microsatellite instability in European hepatocellular Carcinoma", Oncogene, Vol. 18, pp. 181-187, 1999. |

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| | GB | | | | | | |

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| | | |
|--|----|--|
| | GC | Y. Shirota et al., "Identification of Differentially Expressed Genes in Hepatocellular Carcinoma with cDNA Microarrays", Hepatology, Vol. 33, No. 4, pp. 832-840, April 2001. |
| | GD | C. G. Wu et al., "Distinctive gene expression profiles associated with Hepatitis B virus x protein", Oncogene, Vol. 20, pp. 3674-3682, 2001. |
| | GE | L. Xu et al., "Expression Profiling Suggested a Regulatory Role of Liver-Enriched Transcription Factors in Human Hepatocellular Carcinoma", Cancer Research, Vol. 61, pp. 3176-3181, April 1, 2001. |
| | GF | L. Diatchenko et al., "Suppression subtractive hybridization: A method for generating differentially regulated or tissue-specific cDNA probes and libraries", Proc. Natl. Acad. Sci. USA, Vol. 93, pp. 6025-6030, June 1996. |
| | GG | S. F. Altschul et al., "Gapped BLAST and PSI-BLAST: a new generation of protein database search programs", Nucleic Acids Research, Vol. 25, No. 17, pp. 3389-3402, 1997. |
| | GH | J. Brennecke et al., "bantam Encodes a developmentally Regulated microRNA that Controls Cell Proliferation and Regulates the Proapoptotic Gene hid in Drosophila", Cell, Vol. 113, pp. 25-36, April 4, 2003. |
| | GI | H. Kawasaki et al., "Hes1 is a target of microRNA-23 during retinoic-acid-induced neuronal differentiation of NT2 cells", Nature, Vol. 423, pp. 838-842, June 19, 2003. |
| | GJ | J. L. Kissil et al., "Isolation of DAP3, a Novel Mediator of Interferon-γ-induced Cell Death", The Journal of Biological Chemistry, Vol. 270, No. 46, pp. 27932-27936, November 17, 1995. |
| | GK | S. Zimmer et al., "A Novel Human Tocopherol-associated Protein", The Journal of Biological Chemistry, Vol. 275, No. 33, pp. 25672-25680, August 18, 2000. |
| | GL | L. Zhu et al., "Cloning and Characterization of A New Silver-Stainable Protein SSP29, A Member of the LRR Family", Biochemistry and Molecular Biology International, Vol. 42, No. 5, pp. 927-935, August 1997. |

EXAMINER

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|--|----|--------------------|------|---------|-------|----------|-----------------------|
| | HB | | | | | | |

OTHER DOCUMENT(S) (Including Author, Title, Date, Pertinent Pages, Etc.)

| | | |
|--|----|--|
| | HC | C. A. Stein, "Anti-sense Oligodeoxynucleotides - Promises and Pitfalls", Leukemia, Vol. 6, No. 10, pp. 967-974, October 1992. |
| | HD | N. K. Vaish et al., "Recent developments in the hammerhead ribozyme field", Nucleic Acids Research, Vol. 26, No. 23, pp. 5237-5242, 1998. |
| | HE | S. Filleur et al., "SiRNA-mediated Inhibition of Vascular Endothelial Growth Factor Severely Limits Tumor Resistance to Antiangiogenic Thrombospondin-1 and Slows Tumor Vascularization and Growth", Cancer Research, Vol. 63, No. 14, pp. 3919-3922, July 15, 2003. |
| | HF | D. Mumberg et al., "Regulatable promoters of <i>Saccharomyces cerevisiae</i> : comparison of transcription activity and their use for heterologous expression", Nucleic Acids Research, Vol. 22, No. 25, pp. 5767-5768, 1994. |
| | HG | T. R. Brummelkamp et al., "A System for Stable Expression of Short Interfering RNAs in Mammalian Cells", Science, Vol. 296, pp. 550-553, April 19, 2002. |
| | HH | D. W. Russell et al., "Nucleotide Sequence of the Yeast Alcohol Dehydrogenase II Gene", The Journal of Biological Chemistry, Vol. 258, No. 4, pp. 2674-2682, February 25, 1983. |
| | HI | F. Lee et al., "Glucocorticoids regulate expression of dihydrofolate reductase cDNA in mouse mammary tumor virus chimaeric plasmids", Nature, Vol. 294, No. 214, pp. 228-232, November 1981. |
| | HJ | G. Winter et al., "Man-made antibodies", Nature, Vol. 349, pp. 293-299, January 24, 1991. |
| | HK | M. Vidal et al., "Prospects for drug screening using the reverse two-hybrid system", Trends in Biotechnology, Vol. 17, pp. 374-381, September 1999. |
| | HL | P. Fickert et al., "Cytokeratins as Targets for Bile Acid-Induced Toxicity", American Journal of Pathology, Vol. 160, No. 2, pp. 491-499, February 2002. |

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| | IA | | | | | | |

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|--|----|--------------------|------|---------|-------|----------|-----------------------|
| | IB | | | | | | |

OTHER DOCUMENT(S) (Including Author, Title, Date, Pertinent Pages, Etc.)

| | | |
|--|----|--|
| | IC | P. Hedge et al., "A Concise Guide to cDNA Microarray Analysis", Biotechniques, Vol. 29, No. 3, pp. 548-560, 2000. |
| | ID | P. Colas et al., "The impact of two-hybrid and related methods on biotechnology", TIBTECH, Vol. 16, pp. 355-363, August 1998. |
| | IE | A. Knappik et al., "Fully Synthetic Human Combinatorial Antibody Libraries (HuCAL) Based on Modular Consensus Frameworks and CDRs Randomized with Trinucleotides", J. Mol. Biol., Vol. 296, pp. 57-86, 2000. |
| | IF | H. E. Chadd et al., "Therapeutic antibody expression technology", Curr. Opin. Biotechnol., Vol. 12, pp. 188-194, 2001. |
| | IG | P. A. Wood, Transgenic Animal Technology, A Laboratory Handbook, Chapter 5: Retrovirus-Mediated Gene Transfer, pp. 147-176, 1994. |
| | IH | G. M. Monastersky, Transgenic Animal Technology, A Laboratory Handbook, Chapter 6: Gene Transfer Technology: Alternative Techniques and Applications, pp. 177-219, 1994. |
| | II | D. G. Myszkowski et al., "Implementing surface plasmon resonance biosensors in drug discovery", PSTT, Vol. 3, No. 9, pp. 310-317, September 2000. |
| | IJ | P. L. M. Jansen, "Hepatology in the 21 st Century, Gene transfer, hepatocyte transplantation, DNA chips, cyberspace and... and friendly hospital", The Netherlands Journal of Medicine, Vol. 55, pp. 287-292, 1999. |
| | IK | A. A. Khromykh, "Replicon-based vectors of positive strand RNA viruses", Current Opinion in Molecular Therapeutics, Vol. 2, No. 5, pp. 555-569, 2000. |

EXAMINER

DATE CONSIDERED

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INFORMATION DISCLOSURE STATEMENT

FORM PTO 1449 (modified)

U.S. DEPARTMENT OF COMMERCE
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Date Submitted to PTO: July 7, 2005

ATTY DOCKET NO.
2005_0070ASERIAL NO.
10/528,031APPLICANT
Christian GUELLY et al.FILING DATE
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GROUP

U.S. PATENT DOCUMENTS

| *EXAMINER INITIAL | | DOCUMENT NUMBER | DATE | NAME | CLASS | SUBCLASS | FILING DATE IF APPROPRIATE |
|----------------------|----|--------------------|--------|----------------|-------|----------|-------------------------------|
| | JA | 6,899,830 | 5/2005 | Boymond et al. | | | Corresponds to Ref AL |
| | JB | 2003/0157505 | 8/2003 | Miculka et al. | | | Corresponds to Ref AJ |
| | JC | 2002/0028442 | 3/2002 | Miculka et al. | | | Corresponds to Ref AJ |

FOREIGN PATENT DOCUMENTS

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|--|----|--------------------|------|---------|-------|----------|-----------------------|--|
| | JD | | | | | | | |

OTHER DOCUMENT(S) (Including Author, Title, Date, Pertinent Pages, Etc.)

| | | | | | | | | |
|--|----|--|--|--|--|--|--|--|
| | JE | C. R. Graveel et al., "Expression profiling and identification of novel genes in hepatocellular carcinomas", Oncogene, No. 20, pp. 2704-2712, 2001. | | | | | | |
| | JF | M. F. Chen et al., "Partial hepatic resection for hepatocellular carcinoma", Journal of Gastroenterology and Hepatology, Vol. 12, pp. S329-S334, 1997. | | | | | | |
| | JG | R. N. M. MacSween et al., eds., Pathology of the Liver, 4 th Edition, 2001, Churchill Livingstone, Edinburgh, pp. 728-730. | | | | | | |
| | JH | | | | | | | |
| | JI | | | | | | | |
| | JJ | | | | | | | |
| | JK | | | | | | | |

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